UNITED STATES DEPARTMENT OF COMMERCE United States Patent and Trademark Office Address: COMMISSIONER FOR PATENTS P.O. Rey 1459

P O Box 1450 Alexandria, Virgima 22313-1450 www.uspto.gov

NOTICE OF ALLOWANCE AND FEE(S) DUE

26995 7590 022242010 KNOBBE MARTENS OLSON & BEAR LLP 2040 MAIN STREET FOURTEENTH FLOOR

IRVINE, CA 92614

EXAMINER

LEE, JAE W

ART UNIT PAPER NUMBER

1656

DATE MAILED: 02/24/2010

APPLICATION NO.	FILING DATE	FIRST NAMED INVENTOR	ATTORNEY DOCKET NO.	CONFIRMATION NO.	
10/517,225	06/09/2005	Dario Alessi	ERP01.005APC	2433	
TITLE OF INVENTION: METHODS					

APPLN. TYPE	SMALL ENTITY	ISSUE FEE DUE	PUBLICATION FEE DUE	PREV. PAID ISSUE FEE	TOTAL FEE(S) DUE	DATE DUE
nonprovisional	NO	\$1510	\$300	\$0	\$1810	05/24/2010

THE APPLICATION IDENTIFIED ABOVE HAS BEEN EXAMINED AND IS ALLOWED FOR ISSUANCE AS A PATENT, PROSECUTION ON THE MERITS IS CLOSED. THIS NOTICE OF ALLOWANCE IS NOT A GRANT OF PATENT RIGHTS. THIS APPLICATION IS SUBJECT TO WITHDRAWAL FROM ISSUE AT THE INITIATIVE OF THE OFFICE OR UPON PETITION BY THE APPLICANT. SEE 37 CFR 1.313 AND MPEP 1308.

THE ISSUE FEE AND PUBLICATION FEE (IF REQUIRED) MUST BE PAID WITHIN THREE MONTHS FROM THE MAILING DATE OF THIS NOTICE OR THIS APPLICATION SHALL BE REGARDED AS ABANDONED. THIS STATUTORY PERIOD CANNOT BE EXTENDED. SEE 35 U.S.C. 1SI. THE ISSUE FEE DUE INDICATED ABOVE DOES NOT REFLECT A CREDIT FOR ANY PREVIOUSLY PAID ISSUE FEE IN THIS APPLICATION. IF AN ISSUE FEE HAS PREVIOUSLY BEEN PAID IN THIS APPLICATION (AS SHOWN ABOVE), THE RETURN OF PART B OF THIS FORM WILL BE CONSIDERED A REQUEST TO REAPPLY THE PREVIOUSLY PAID ISSUE FEE TOWARD THE ISSUE FEE NOW DUE.

HOW TO REPLY TO THIS NOTICE:

I. Review the SMALL ENTITY status shown above.

If the SMALL ENTITY is shown as YES, verify your current SMALL ENTITY status:

A. If the status is the same, pay the TOTAL FEE(S) DUE shown above.

B. If the status above is to be removed, check box 5b on Part B - Fee(s) Transmittal and pay the PUBLICATION FEE (if required) and twice the amount of the ISSUE FEE shown above, or

If the SMALL ENTITY is shown as NO:

A. Pay TOTAL FEE(S) DUE shown above, or

B. If applicant claimed SMALL ENTITY status before, or is now claiming SMALL ENTITY status, check box 5a on Part B - Fee(s) Transmittal and pay the PUBLICATION FEE (if required) and 1/2 the ISSUE FIEE shown above.

II. PART B - FEE(S) TRANSMITTAL, or its equivalent, must be completed and returned to the United States Patent and Trademark Office (USPTO) with your ISSUE FEE and PUBLICATION FEE (if required). If you are charging the fee(s) to your deposit account, section "4b" of Part B - Fee(s) Transmittal should be completed and an extra copy of the form should be submitted. If an equivalent of Part B is filed, a request to reapply a previously paid issue fee must be clearly made, and delays in processing may occur due to the difficulty in recognizing the paper as an equivalent of Part B.

III. All communications regarding this application must give the application number. Please direct all communications prior to issuance to Mail Stop ISSUE FEE unless advised to the contrary.

IMPORTANT REMINDER: Utility patents issuing on applications filed on or after Dec. 12, 1980 may require payment of maintenance fees. It is patentee's responsibility to ensure timely payment of maintenance fees when due.

PART B - FEE(S) TRANSMITTAL

Complete and send this form, together with applicable fee(s), to: Mail Commissioner for Patents P.O. Box 1450 Alexandria, Virginia 22313-1450

or Fax (571)-273-2885

INSTRUCTIONS: This appropriate. All further a indicated unless correcte maintenance fee notificat	a below or directed our	or transmitting the ISSU ig the Patent, advance on nerwise in Block 1, by (a	JE FEE and PUBLIC rders and notification a) specifying a new o	orrespondence addres	s; and/o	or (b) indicating a sepa	hould be completed where correspondence address as arate "FEE ADDRESS" for
CURRENT CORRESPONDENCE ADDRESS (Note: Use Block 1 for any change of address)				Note: A certificate of mailing can only be used for domestic mailings of the Fe(s) Transmittal. This certificate cannot be used for any other accompanying papers. Each additional paper, such as an assignment or formal drawing, must have its own certificate of mailing or transmission.			
20995	7590 02/24	/2010					
KNOBBE MAI 2040 MAIN STR FOURTEENTH IRVINE, CA 926	FLOOR	& BEAR LLP		I hereby certify that States Postal Service addressed to the Ma transmitted to the US	ertificat this Feet with su til Stop PTO (57	e of Mailing or Trans (s) Transmittal is being fficient postage for fir ISSUE FEE address 71) 273-2885, on the d	mission g deposited with the United st class mail in an envelope above, or being facsimile ate indicated below.
IR VINE, CA 920	014						(Depositor's name)
							(Signature)
							(Date)
APPLICATION NO.	FILING DATE		FIRST NAMED INVEN	TOR	ATTO	ORNEY DOCKET NO.	CONFIRMATION NO.
10/517,225	06/09/2005	•	Dario Alessi			ERP01.005APC	2433
TITLE OF INVENTION:	METHODS						
APPLN. TYPE	SMALL ENTITY	ISSUE FEE DUE	PUBLICATION FEE D	UE PREV. PAID ISS	UE FEE	TOTAL FEE(S) DUE	DATE DUE
nonprovisional	NO	\$1510	\$300	\$0		\$1810	05/24/2010
EXAM	INER	ART UNIT	CLASS-SUBCLASS				
LEE, J.		1656	703-011000				
"Fee Address" indi PTO/SB/47; Rev 03-0: Number is required. 3. ASSIGNEE NAME AT	ondence address (or Cha 1/122) attached. cation (or "Fee Address' 2 or more recent) attach ND RESIDENCE DATA ess an assignee is identia n in 37 CFR 3.11. Comp	nge of Correspondence Indication form and. Use of a Customer A TO BE PRINTED ON	(I) the names of u or agents OR, alter (2) the name of a sregistered attorney 2 registered patent listed, no name will THE PATENT (print of data will appear on the or a substitute for filing the or agent of	ingle firm (having as or agent) and the na attorneys or agents. I I be printed.	a memi mes of u f no nar	ber a 2	ocument has been filed for
Please check the appropri	ate assignee category or	categories (will not be pr	rinted on the patent):	☐ Individual ☐ 6	Corporat	tion or other private gr	oup entity Government
4a. The following fee(s) are submitted: Issue Fee Publication Fee (No small entity discount permitted) Advance Order - # of Copies			A check is enclos Payment by credi	ed. t card. Form PTO-20:	38 is att	required fee(s), any de	
	SMALL ENTITY statu	is. See 37 CFR 1.27.				TITY status. Sec 37 C	
NOTE: The Issue Fee and interest as shown by the r	d Publication Fee (if requeecords of the United Sta	uired) will not be accepte tes Patent and Trademark	d from anyone other the Office.	an the applicant; a re	gistered	attorney or agent; or the	ne assignee or other party in
Authorized Signature				Date			
Typed or printed name				Registration No.			
This collection of informa an application. Confident submitting the completed this form and/or suggestic Box 1450, Alexandria, V Alexandria, Virginia 223	ation is required by 37 C iality is governed by 35 application form to the ons for reducing this but irginia 22313-1450. DC 13-1450.	FR 1.311. The informatic U.S.C. 122 and 37 CFR USPTO. Time will vary rden, should be sent to the ONOT SEND FEES OR	on is required to obtain 1.14. This collection is depending upon the e Chief Information O COMPLETED FORM	or retain a benefit by s estimated to take 12 ndividual case. Any fficer, U.S. Patent an S TO THIS ADDRE:	the pub minute commen d Trader SS. SEN	olic which is to file (and is to complete, including the on the amount of timark Office, U.S. Dep ID TO: Commissioner	d by the USPTO to process) ig gathering, preparing, and me you require to complete artment of Commerce, P.O. for Patents, P.O. Box 1450,

Under the Paperwork Reduction Act of 1995, no persons are required to respond to a collection of information unless it displays a valid OMB control number.



UNITED STATES PATENT AND TRADEMARK OFFICE

UNITED STATES DEPARTMENT OF COMMERCE United States Patent and Trademark Office Address: COMMISSIONER FOR PATENTS

P O Box 1450 Alexandria, Virgima 22313-1450 www.uspto.gov

APPLICATION NO.	FILING DATE	FIRST NAMED INVENTOR	ATTORNEY DOCKET NO.	CONFIRMATION NO.	
10/517,225	06/09/2005	Dario Alessi	ERP01.005APC	2433	
20995 75	90 02/24/2010		EXAM	UNER	
KNOBBE MAR	TENS OLSON & BE	LEE, JAE W			
2040 MAIN STRE			ART UNIT	PAPER NUMBER	
FOURTEENTH FLOOR IRVINE, CA 92614			1656		
IIX VIIVE, CA 5201	*	DATE MAIL ED: 02/24/2010			

Determination of Patent Term Adjustment under 35 U.S.C. 154 (b)

(application filed on or after May 29, 2000)

The Patent Term Adjustment to date is 974 day(s). If the issue fee is paid on the date that is three months after the mailing date of this notice and the patent issues on the Tuesday before the date that is 28 weeks (six and a half months) after the mailing date of this notice, the Patent Term Adjustment will be 974 day(s).

If a Continued Prosecution Application (CPA) was filed in the above-identified application, the filing date that determines Patent Term Adjustment is the filing date of the most recent CPA.

Applicant will be able to obtain more detailed information by accessing the Patent Application Information Retrieval (PAIR) WEB site (http://pair.uspto.gov).

Any questions regarding the Patent Term Extension or Adjustment determination should be directed to the Office of Patent Legal Administration at (571)-272-7702. Questions relating to issue and publication fee payments should be directed to the Customer Service Center of the Office of Patent Publication at 1-(888)-786-0101 or (571)-272-4200.

Application No. Applicant(s) 10/517.225 ALESSI ET AL. Notice of Allowability Examiner Art Unit JAE W. LEE 1656 -- The MAILING DATE of this communication appears on the cover sheet with the correspondence address--All claims being allowable, PROSECUTION ON THE MERITS IS (OR REMAINS) CLOSED in this application. If not included herewith (or previously mailed), a Notice of Allowance (PTOL-85) or other appropriate communication will be mailed in due course. THIS NOTICE OF ALLOWABILITY IS NOT A GRANT OF PATENT RIGHTS. This application is subject to withdrawal from issue at the initiative of the Office or upon petition by the applicant. See 37 CFR 1.313 and MPEP 1308. This communication is responsive to 01/18/2010. The allowed claim(s) is/are 1-8,11-13,16,43 and 44. 3. Acknowledgment is made of a claim for foreign priority under 35 U.S.C. § 119(a)-(d) or (f). a) 🔯 All b) ☐ Some* c) ☐ None of the: 1. T Certified copies of the priority documents have been received. 2. Certified copies of the priority documents have been received in Application No. 3. \(\overline{\text{ Copies of the certified copies of the priority documents have been received in this national stage application from the International Bureau (PCT Rule 17.2(a)). * Certified copies not received: _____. Applicant has THREE MONTHS FROM THE "MAILING DATE" of this communication to file a reply complying with the requirements noted below. Failure to timely comply will result in ABANDONMENT of this application. THIS THREE-MONTH PERIOD IS NOT EXTENDABLE. A SUBSTITUTE OATH OR DECLARATION must be submitted. Note the attached EXAMINER'S AMENDMENT or NOTICE OF INFORMAL PATENT APPLICATION (PTO-152) which gives reason(s) why the oath or declaration is deficient. CORRECTED DRAWINGS (as "replacement sheets") must be submitted. (a) Including changes required by the Notice of Draftsperson's Patent Drawing Review (PTO-948) attached 1) hereto or 2) to Paper No./Mail Date (b) including changes required by the attached Examiner's Amendment / Comment or in the Office action of Identifying indicia such as the application number (see 37 CFR 1.84(c)) should be written on the drawings in the front (not the back) of each sheet. Replacement sheet(s) should be labeled as such in the header according to 37 CFR 1.121(d). 6. DEPOSIT OF and/or INFORMATION about the deposit of BIOLOGICAL MATERIAL must be submitted. Note the attached Examiner's comment regarding REQUIREMENT FOR THE DEPOSIT OF BIOLOGICAL MATERIAL. Attachment(s) 1. | Notice of References Cited (PTO-892) 5. Notice of Informal Patent Application 2. Notice of Draftperson's Patent Drawing Review (PTO-948) Interview Summary (PTO-413), Paper No./Mail Date Information Disclosure Statements (PTO/SB/08). 7. X Examiner's Amendment/Comment

Paper No./Mail Date

of Biological Material

4. T Examiner's Comment Regarding Requirement for Deposit

8. T Examiner's Statement of Reasons for Allowance

Other .

Art Unit: 1656

EXAMINER'S AMENDMENT

An examiner's amendment to the record appears below. Should the changes and/or additions be unacceptable to applicant, an amendment may be filed as provided by 37 CFR 1.312. To ensure consideration of such an amendment, it MUST be submitted no later than the payment of the issue fee.

Authorization for this Examiner's amendment was given in a telephone interview with Daniel Altman on 02/10/2010. Claims 1-8, 11-13, 16, 43 and 44 are allowed.

The title of the invention is not descriptive. A new title is required that is clearly indicative of the invention to which the claims are directed.

Please replace the title "Methods" with "Methods for selecting a candidate compound using a three-dimensional structure of phosphoinositide dependent protein kinase 1 (PDK1)".

Please amend the claims as they appear below:

A clean copy of Claim 1 is provided as shown below:

Claim 1. A method for selecting a compound for modulating the activity of phosphoinositide dependent protein kinase 1 (PDK1), the method comprising:

Art Unit: 1656

[i] generating a three-dimensional structure of the protein kinase catalytic domain of PDK1 as defined by Example 2, 3, 7 or 8;

[ii] identifying predicted binding region(s): the PDK1 Interacting Fragment (PIF) binding pocket formed by residues including residues Lys115, Ile118, Ile119 on the α B helix, Val124, Val127 on the α C helix, and Leu 155 on β -sheet 5 of full length human PDK1 as set forth in SEQ ID NO: 3, the "phosphate binding pocket" formed by residues including residues Lys76, Arg131, Thr148 and Gln150 of full length human PDK1 as set forth in SEQ ID NO: 3, and/or the α C helix formed by residues 123-136 of full length human PDK1 as set forth in SEQ ID NO: 3 in any of the models generated in step [i];

- [iii] modeling a plurality of three-dimensional structures of candidate compounds;
- [iv] comparing said three-dimensional structure of the protein kinase catalytic domain of PDK1 binding regions/pockets as identified in step [ii] with the three-dimensional structure of each of said candidate compounds of step [iii]; and
- [v] selecting one of the candidate compounds as a compound for modulating the activity of phosphoinositide dependent protein kinase 1 (PDK1).

Art Unit: 1656

Claim 2. Replace "(SEQ ID NO. 3) or a fusion thereof" with "as set forth in SEQ ID NO: 3".

Claim 3. Replace "claim 2" in line 1 with "claim 1"; and replace "(SEQ ID NO. 3)" with "as set forth in SEQ ID NO: 3".

Claim 4. Replace "(SEQ ID NO. 3) or a fusion thereof" with "as set forth in SEQ ID NO: 3".

Claims 9 and 10. Canceled.

Claim 12. Replace "(SEQ ID NO: 2)" with "as set forth in SEQ ID NO: 2".

A clean copy of Claim 13 is provided as shown below:

Claim 13. A method for selecting a compound for modulating the activity of PDK1 Interacting Fragment (PIF) binding pocket containing protein kinase, the method comprising:

[i] modeling a three-dimensional structure of a plurality of molecules in a computer;

Art Unit: 1656

[ii] generating a three-dimensional structure of the protein kinase catalytic domain of PDK1 as defined by Example 2, 3, 7 or 8;

[iii] comparing the three-dimensional structure of each of said molecules with the three-dimensional structure of said phosphate binding pocket of human PDK1 and optionally also said hydrophobic pocket of human PDK1:

[iv] identifying a molecule from the plurality of the modeled molecules which binds to said PIF binding pocket in the position equivalent to a hydrophobic pocket of human PDK1 that is defined by residues including Lys115, Ile118, Ile119, Val124, Val127 and Leu155 of full-length human PDK1 as set forth in SEQ ID NO: 3, and/or a phosphate binding pocket in the position equivalent to the phosphate binding pocket of human PDK1 that is defined by residues including Lys76, Arg131, Thr148 and Gln150 of SEQ ID NO: 3; and

[v] selecting one of the modeled molecules as said compound based on a predicted ability of the molecule to interact with said phosphate binding pocket of protein kinase or said hydrophobic pocket of protein kinase,

Art Unit: 1656

wherein the three-dimensional structure or a part thereof is determined for a polypeptide comprising residues 51 to 359 of full length human PDK1 as set forth in SEQ ID NO: 3.

Claims 14 and 15. Canceled.

Claims 43 and 44. Replace "prior" with "prior to".

Any comments considered necessary by applicant must be submitted no later than the payment of the issue fee and, to avoid processing delays, should preferably accompany the issue fee. Such submissions should be clearly labeled "Comments on Statement of Reasons for Allowance."

Any inquiry concerning this communication or earlier communications from the examiner should be directed to Jae W. Lee whose telephone number is 571-272-9949. The examiner can normally be reached between 9:00 to 5:30 on Monday-Friday.

If attempts to reach the examiner by telephone are unsuccessful, the examiner's supervisor, Manjunath Rao can be reached on 571-272-0939. The fax phone number for the organization where this application or proceeding is assigned is 571-273-8300.

Information regarding the status of an application may be obtained from the Application Information Retrieval (PAIR) system. Status information for published applications may be obtained from either Private PAIR or Public PAIR. Status information for unpublished applications is available through Private PAIR only. For

Art Unit: 1656

more information about the PAIR system, see http://pair-direct.uspto.gov. Should you have questions on access to the Private PAIR system, contact the Electronic Business Center (EBC) at 866-217-9197 (toll-free). If you would like assistance from a USPTO Customer Service Representative or access to the automated information system, call 800-786-9199 (IN USA OR CANADA) or 571-272-1000.

/JAE W LEE/ Examiner, Art Unit 1656

/SUZANNE M. NOAKES/ Primary Examiner, Art Unit 1656